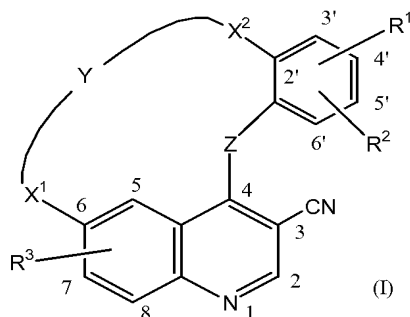


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ABSTRACT

3-CYANO-QUINOLINE DERIVATIVES WITH ANTIPROLIFERATIVE ACTIVITY

5 The present invention concerns the compounds of formula



the *N*-oxide forms, the pharmaceutically acceptable addition salts and the stereochemically isomeric forms thereof, wherein

10 Z represents NH;

Y represents -C₃₋₉alkyl-, -C₁₋₅alkyl-NR¹²-C₁₋₅alkyl-, -C₁₋₆alkyl-NH-CO- or
-CO-NH -C₁₋₆alkyl- ;

X¹ represents -O-;

15 X² represents a direct bond, -NR¹¹-C₁₋₂alkyl-, -NR¹¹-CH₂-, -C₁₋₂alkyl-, -O-C₁₋₂alkyl,
-O- or -O-CH₂-;

R¹ represents hydrogen or halo;

R² represents hydrogen, cyano, halo, hydroxycarbonyl-, C₁₋₄alkyloxycarbonyl-,
Het¹⁶-carbonyl- or Ar⁵;

20 R³ represents hydrogen, hydroxy, C₁₋₄alkyloxy-, Ar⁴-C₁₋₄alkyloxy or R³ represents
C₁₋₄alkyloxy substituted with one or where possible two or more substituents
selected from C₁₋₄alkyloxy- or Het²-;

R¹⁰ represents hydrogen;

R¹¹ represents hydrogen, C₁₋₄alkyl- or C₁₋₄alkyl-oxy-carbonyl-;

R¹² represents Het¹⁴-C₁₋₄alkyl-, in particular morpholinyl-C₁₋₄alkyl-;

25 Het² represents a heterocycle selected from morpholinyl or piperidinyl optionally
substituted with C₁₋₄alkyl-, preferably methyl;

Het¹⁴ represents morpholinyl;

Het¹⁶ represents a heterocycle selected from morpholinyl or pyrrolidinyl;

Ar⁴ represents phenyl;

30 Ar⁵ represents phenyl optionally substituted with cyano.